

FORM PTO-1390 (REV. 9-2001)		U.S. DEPARTMENT OF COMMERCE PATENT AND TRADEMARK OFFICE		ATTORNEY'S DOCKET NUMBER 366325-525	
TRANSMITTAL LETTER TO THE UNITED STATES DESIGNATED/ELECTED OFFICE (DO/EO/US) CONCERNING A FILING UNDER 35 U.S.C. 371				U.S. APPLICATION NO. (if known, see 37 CFR 1.5)	
				10/088829 Not yet assigned	
INTERNATIONAL APPLICATION NO. PCT/FR00/02668		INTERNATIONAL FILING DATE 27 September 2000 (27.09.2000)		PRIORITY DATE CLAIMED 27 September 1999 (27.09.1999)	
TITLE OF INVENTION Method and Installation For Setting In Adsorbed Stated On A Porous Support Active Compounds Contained in					
APPLICANT(S) FOR DO/EO/US Perrut, et al.					
Applicant herewith submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information:					
<p>1. <input checked="" type="checkbox"/> This is a FIRST submission of items concerning a filing under 35 U.S.C. 371.</p> <p>2. <input type="checkbox"/> This is a SECOND or SUBSEQUENT submission of items concerning a filing under 35 U.S.C. 371.</p> <p>3. <input checked="" type="checkbox"/> This is an express request to begin national examination procedures (35 U.S.C. 371(f)). The submission must include items (5), (6), (9) and (21) indicated below.</p> <p>4. <input type="checkbox"/> The US has been elected by the expiration of 19 months from the priority date (Article 31).</p> <p>5. <input checked="" type="checkbox"/> A copy of the International Application as filed (35 U.S.C. 371(c)(2))</p> <p>a. <input checked="" type="checkbox"/> is attached hereto (required only if not communicated by the International Bureau).</p> <p>b. <input type="checkbox"/> has been communicated by the International Bureau.</p> <p>c. <input type="checkbox"/> is not required, as the application was filed in the United States Receiving Office (RO/US).</p> <p>6. <input checked="" type="checkbox"/> An English language translation of the International Application as filed (35 U.S.C. 371(c)(2)).</p> <p>a. <input checked="" type="checkbox"/> is attached hereto.</p> <p>b. <input type="checkbox"/> has been previously submitted under 35 U.S.C. 154(d)(4).</p> <p>7. <input type="checkbox"/> Amendments to the claims of the International Application under PCT Article 19 (35 U.S.C. 371(c)(3))</p> <p>a. <input type="checkbox"/> are attached hereto (required only if not communicated by the International Bureau).</p> <p>b. <input type="checkbox"/> have been communicated by the International Bureau.</p> <p>c. <input type="checkbox"/> have not been made; however, the time limit for making such amendments has NOT expired.</p> <p>d. <input type="checkbox"/> have not been made and will not be made.</p> <p>8. <input type="checkbox"/> An English language translation of the amendments to the claims under PCT Article 19 (35 U.S.C. 371 (c)(3)).</p> <p>9. <input checked="" type="checkbox"/> An oath or declaration of the inventor(s) (35 U.S.C. 371(c)(4)).</p> <p>10. <input checked="" type="checkbox"/> An English language translation of the annexes of the International Preliminary Examination Report under PCT Article 36 (35 U.S.C. 371(c)(5)).</p>					
Items 11 to 20 below concern document(s) or information included:					
<p>11. <input checked="" type="checkbox"/> An Information Disclosure Statement under 37 CFR 1.97 and 1.98.</p> <p>12. <input checked="" type="checkbox"/> An assignment document for recording. A separate cover sheet in compliance with 37 CFR 3.28 and 3.31 is included.</p> <p>13. <input type="checkbox"/> A FIRST preliminary amendment.</p> <p>14. <input type="checkbox"/> A SECOND or SUBSEQUENT preliminary amendment.</p> <p>15. <input type="checkbox"/> A substitute specification.</p> <p>16. <input checked="" type="checkbox"/> A change of power of attorney and/or address letter.</p> <p>17. <input type="checkbox"/> A computer-readable form of the sequence listing in accordance with PCT Rule 13ter.2 and 35 U.S.C. 1.821 - 1.825.</p> <p>18. <input type="checkbox"/> A second copy of the published international application under 35 U.S.C. 154(d)(4).</p> <p>19. <input type="checkbox"/> A second copy of the English language translation of the international application under 35 U.S.C. 154(d)(4).</p> <p>20. <input checked="" type="checkbox"/> Other items or information: Verification of English language translation and acknowledgement postcard</p>					

U.S. APPLICATION NO. (if known, see 37 CFR 1.55) Not yet assigned 108829		INTERNATIONAL APPLICATION NO. PCT/FR00/02668		ATTORNEY'S DOCKET NUMBER 366325-525	
--	--	--	--	---	--

21. <input checked="" type="checkbox"/> The following fees are submitted: BASIC NATIONAL FEE (37 CFR 1.492 (a) (1) - (5)): Neither international preliminary examination fee (37 CFR 1.482) nor international search fee (37 CFR 1.445(a)(2)) paid to USPTO and International Search Report not prepared by the EPO or JPO \$1040.00 International preliminary examination fee (37 CFR 1.482) not paid to USPTO but International Search Report prepared by the EPO or JPO \$890.00 International preliminary examination fee (37 CFR 1.482) not paid to USPTO but international search fee (37 CFR 1.445(a)(2)) paid to USPTO \$740.00 International preliminary examination fee (37 CFR 1.482) paid to USPTO but all claims did not satisfy provisions of PCT Article 33(1)-(4) \$710.00 International preliminary examination fee (37 CFR 1.482) paid to USPTO and all claims satisfied provisions of PCT Article 33(1)-(4) \$100.00 ENTER APPROPRIATE BASIC FEE AMOUNT =				CALCULATIONS PTO USE ONLY <table style="width:100%; border-collapse: collapse;"> <tr> <td style="width:50%; text-align: right;">\$</td> <td style="width:50%; text-align: right;">890.00</td> </tr> <tr> <td style="text-align: right;">\$</td> <td style="text-align: right;">0.00</td> </tr> </table>		\$	890.00	\$	0.00
\$	890.00								
\$	0.00								
Surcharge of \$130.00 for furnishing the oath or declaration later than <input type="checkbox"/> 20 <input type="checkbox"/> 30 months from the earliest claimed priority date (37 CFR 1.492(e)).				<table style="width:100%; border-collapse: collapse;"> <tr> <td style="width:50%; text-align: right;">\$</td> <td style="width:50%; text-align: right;">0.00</td> </tr> </table>		\$	0.00		
\$	0.00								
CLAIMS	NUMBER FILED	NUMBER EXTRA	RATE						
Total claims	40 - 20 =	20	x \$18.00	\$	360.00				
Independent claims	2 - 3 =	0	x \$84.00	\$	0.00				
MULTIPLE DEPENDENT CLAIM(S) (if applicable)			+ \$282.00	\$	282.00				
TOTAL OF ABOVE CALCULATIONS =				\$	1,532.00				
<input type="checkbox"/> Applicant claims small entity status. See 37 CFR 1.27. The fees indicated above are reduced by 1/2.				\$	0.00				
SUBTOTAL =				\$					
Processing fee of \$130.00 for furnishing the English translation later than <input type="checkbox"/> 20 <input type="checkbox"/> 30 months from the earliest claimed priority date (37 CFR 1.492(f)).				\$	0.00				
TOTAL NATIONAL FEE =				\$	1,532.00				
Fee for recording the enclosed assignment (37 CFR 1.21(h)). The assignment must be accompanied by an appropriate cover sheet (37 CFR 3.28, 3.31). \$40.00 per property +				\$	40.00				
TOTAL FEES ENCLOSED =				\$	1,572.00				
				Amount to be refunded:	\$				
				charged:	\$				

a. ☒ A check in the amount of \$ 1,572.00 to cover the above fees is enclosed.

b. ☐ Please charge my Deposit Account No. _____ in the amount of \$ _____ to cover the above fees.
 A duplicate copy of this sheet is enclosed.

c. ☒ The Commissioner is hereby authorized to charge any additional fees which may be required, or credit any
 overpayment to Deposit Account No. 04-0480. A duplicate copy of this sheet is enclosed.

d. ☐ Fees are to be charged to a credit card. **WARNING:** Information on this form may become public. **Credit card
 information should not be included on this form.** Provide credit card information and authorization on PTO-2038.

NOTE: Where an appropriate time limit under 37 CFR 1.494 or 1.495 has not been met, a petition to revive (37 CFR
 1.137 (a) or (b)) must be filed and granted to restore the application to pending status.

SEND ALL CORRESPONDENCE TO:

Thomas S. Deibert
 c/o Dechert
 Princeton Pike Corporate Center
 P.O. Box 5218
 Princeton, NJ 08543-5218

SIGNATURE
 Thomas S. Deibert
 NAME
 40,984
 REGISTRATION NUMBER

1008829 PCT/FR00/02668

METHOD AND INSTALLATION FOR SETTING IN ADSORBED STATE
ON A POROUS SUPPORT ACTIVE COMPOUNDS CONTAINED IN A
PRODUCT

The present invention relates to a method and an installation allowing,
5 industrially, the setting in adsorbed state, on a porous support, of certain active
compounds contained in natural or synthetic products. More precisely, such an
operation will be effected after a preliminary phase during which said active
compounds will be extracted from these products with the aid of a solvent taken
to supercritical pressure, i.e. a fluid in supercritical state or a subcritical liquid.

10 In effect, it is known that bodies are generally known in three states,
namely solid, liquid or gaseous, and that one passes from one to the other by
varying the temperature and/or the pressure. Now, there is a point beyond which
one can pass from the liquid state to the gaseous or vapour state without passing
via a boiling or, inversely, a condensation, but continuously: this point is called
15 the critical point.

It is also known that a fluid in supercritical state, i.e. a fluid which is in a
state characterized either by a pressure and a temperature respectively greater
than the critical pressure and temperature in the case of a pure body, or by a
representative point (pressure, temperature) located beyond the envelope of the
20 critical points shown in a diagram (pressure, temperature) in the case of a
mixture, presents, for very numerous substances, a high solvent power without
comparison with that observed in this same fluid

in the state of compressed gas. The same applies to the so-called "subcritical" liquids, i.e. liquids which are in a state characterized either by a pressure higher than the critical pressure and by a temperature lower than the critical temperature in the case of a pure body, or by a pressure higher than the critical pressures and a temperature lower than the critical temperatures of the components in the case of a mixture (cf. on this subject the article by Michel PERRUT - Les Techniques de l'Ingénieur "Extraction by supercritical fluid, J 2 770 - 1 to 12, 1999").

The considerable and modulatable variations of the solvent power of the supercritical fluids are, moreover, used in numerous methods of extraction (solid/fluid), fractionation (liquid/fluid), analytical or preparative chromatography, treatment of materials (ceramics, polymers...). Chemical or biochemical reactions are also made in such solvents. It should be noted that the physico-chemical properties of carbon dioxide as well as its critical parameters (critical pressure: 7.4 MPa and critical temperature: 31°C) make of it the preferred solvent in numerous applications, all the more so as it does not present any toxicity and is available at very low price in very large quantities. Non-polar solvent, the carbon dioxide taken to supercritical pressure sometimes has a co-solvent added thereto, constituted in particular by a polar organic solvent whose function is to considerably modify the solvent power, especially with respect to molecules presenting a certain polarity, ethanol often being used to

that purpose. However, certain compounds are more favourably extracted with a light hydrocarbon having from 2 to 5 carbon atoms, and more favourably, from 2 to 4 carbon atoms, at supercritical pressure.

One of the principal advantages of the methods using fluids at supercritical pressure as solvents resides in the facility of effecting the separation between the solvent and the extracts and solutes, as has been described in numerous publications and, for certain important aspects of implementation, in French Patent FR-A-2 584 618. The interesting properties of these fluids have, moreover, been used for a long time in solid-fluid extraction and liquid-fluid fractionation, as has been described in the article mentioned above.

In the event of the extracts or solutes comprising very volatile products, such as odorant products, the mere separation by partial decompression bringing about separation of the solutes and their collection by phase separation with the fluid, is not very efficient and the fluid, even partially decompressed, still contains substantial quantities of such substances. In order to separate these substances from the fluid before its recycling, it is known to use an adsorbent bed which will fix these substances and purify the fluid, as described for example in Japanese Patent JP-A-02139003.

Finally, it is known that the extraction of natural products by a fluid at supercritical pressure leads to extracts of very high quality which are increasingly used in numerous applications. However, such extracts, like, moreover, the extracts obtained with

other means such as for example extraction by organic solvent, are often in the form of very viscous or even pasty products which are not easy to handle, with the result that their incorporation in solid supports, the dosage and the mixture with a matrix and possibly other active principles, within a solid excipient, are very difficult. One is sometimes obliged to place them in solution in an organic solvent in order to effect impregnation of a solid excipient, which is regrettable since there thus disappears a determining advantage in numerous applications for

AMENDED SHEET

which any contact of the product with an organic solvent is to be avoided.

The present invention has for its object to propose means making it possible, for purposes of industrial production, to extract active principles, particularly of pharmaceutical, cosmetological, dietetic interest, from diverse raw materials in which they are diluted, in variable concentration, depending on the origin of these raw materials and the period of their harvesting, as is always the case for products of natural origin, and to fix the extract obtained in the course of extraction in an adequate porous matrix by impregnation in one and the same operation.

According to the invention, the operation of extraction by fluid at supercritical pressure is coupled with a second operation during which the separation of the extract mixed with the solvent fluid and the impregnation of a porous medium by this extract are effected simultaneously.

The present invention thus has for its object a method for setting in adsorbed state, on a porous support, compounds contained in a product, in which, during a first step, the extraction of the compounds is effected by contacting the product with at least one solvent fluid at supercritical pressure leading to the obtaining of a mixture of extracts and of solvent fluid, characterized in that, in a second step, the water contained in the mixture of extracts and of solvent fluid is eliminated, the temperature and pressure conditions are adjusted so as to obtain, in an enclosure, two phases, namely a

first phase essentially constituted by the solvent fluid in the gaseous state and a second phase constituted by a mixture of liquids formed by solvent fluid and extracts of the product, these two phases are made to trickle through a porous support adapted to adsorb the extracts, the solvent fluid contained in the second
5 phase is vaporized.

The elimination of the water will preferably be ensured by causing the mixture of extracts and of solvent fluid to trickle over a bed of adsorbent product adapted to fix the water selectively.

The solvent fluid may be constituted by pure carbon dioxide, nitrogen
10 protoxide or a light hydrocarbon counting from 2 to 4 carbon atoms. The solvent fluid may be pure or possibly have one or more co-solvents added thereto. For example, the solvent fluid may in particular be constituted by a mixture of carbon dioxide with at least one co-solvent constituted by an alcohol and preferably ethanol, by a ketone and preferably acetone, by an ester and preferably ethyl
15 acetate.

The first step of extraction may preferably be effected at a pressure included between 7.4 MPa and 80 Mpa, and preferably between 10 MPa and 40 MPa, and at a temperature included between 0°C and 80°C. Similarly, the trickling of the two phases through the porous support may be effected at a
20 pressure included between 1 MPa and 10 MPa, and preferably between 4 MPa and 8 Mpa, and at a temperature included between 0°C and 80°C.

The present invention also has for its object an installation for extraction/impregnation of the type comprising an extractor containing a product from which it is desired

to extract the compounds, which is traversed to that end by at least one solvent fluid at supercritical pressure, characterized in that it successively comprises, downstream of the extractor, means for eliminating the water contained in the extracted compounds, means adapted to create, in an impregnation enclosure
5 containing a porous medium, two phases, namely a first phase essentially constituted by the solvent fluid in the gaseous state and a second phase constituted by a mixture of liquids formed by solvent fluid and the extracts of the product, so as to effect the adsorption by the porous medium of the extracted compounds. The impregnation enclosure may comprise means for contributing
10 enthalpy, particularly constituted by a double envelope with circulation of heat-exchange fluid.

The installation may also comprise, downstream of the impregnation enclosure, means for condensation of the solvent fluid.

In a variant embodiment of the invention, the extractor may be constituted
15 by a fractionating column operating in counter-flow, adapted for the treatment of liquid raw materials.

Various forms of embodiment of the present invention will be described hereinafter by way of non-limiting examples, with reference to the accompanying single Figure which schematically shows an installation for carrying out the
20 method according to the invention.

As has been described in detail in the article and the Patent mentioned hereinabove, it is known that the methods of extraction and of fractionation using a fluid at supercritical pressure comprise two successive steps:

a first step during which the solvent fluid is contacted with the raw material to be treated, under such pressure and temperature conditions that its solvent power with respect to the compounds to be extracted is high, and a second step during which the fluid is placed under such pressure and temperature conditions that its solvent power is very weak with respect to the products which it has extracted from the raw material during the earlier step, which allows the separation of these extracts, the fluid then being recycled. Unlike the conventional embodiments consisting in separating the two phases obtained during this second step in gravity or inertia separators, the method according to the invention consists in causing the mixture of these two phases to trickle within a porous medium.

It has been noticed that, if the operation is carried out without particular precaution, the extracts are not distributed homogeneously on and within the porous medium and, even by proceeding with stirring of this porous medium if it is in pulverulent or granular form, for example with the aid of a rotating turbine, lumps of variable size are generally obtained, resulting from the agglomeration of the particles of the porous medium by the extracts which, instead of penetrating in the pores, remain on the surface without homogeneous distribution. Moreover, natural products, capable of being advantageously treated by a method of this type, always having a certain content of water, the extracts always contain water of which the presence complicates their adsorption on the porous supports or excipients, as the majority of them have a great

affinity for water, so that their structure and their mechanical properties are seriously altered by the water thus contributed by the fluid at the same time as the extracts envisaged.

On the contrary, when the water is trapped, particularly by a selective
5 adsorbent, and the impregnation of the porous medium is then effected under determined pressure and temperature conditions, it is observed that, surprisingly, the extracts impregnate this porous medium in very homogeneous and reproducible manner, leading for example to a non-sticky powder presenting a good fluidity in the case of the initial porous medium also presenting these
10 characteristics.

An example of installation for carrying out the method according to the invention will be described hereinafter with reference to the single Figure.

This installation is derived from a conventional unit of extraction by fluid at supercritical pressure intended for the discontinuous treatment of solid materials.

15 It comprises an extractor 1 containing a basket 2 intended to receive the raw material to be treated, and a diaphragm pump 3 which distributes liquid carbon dioxide at the working pressure, through a heat exchanger 4 making it possible to heat the fluid to the working temperature. Unlike the conventional installations for extraction by fluid at supercritical pressure, the fluid emerging from the extractor
20 1 is conducted in a recipient 5 which contains a porous medium for selectively adsorbing water, such as in particular the molecular sieve 3A. The fluid which is under the same pressure and temperature conditions

as those prevailing in the extractor 1, trickles through the porous medium where it abandons the water that it contains.

5 The recipient 5 is connected to a pressure reducing valve 8 via a heater-exchanger 7. The outlet of the valve 8 is connected to the bottom of an impregnation enclosure 9 which contains the porous medium 10 in which it is desired to adsorb the extracts. The enclosure 9 comprises heating means constituted for example by a double envelope 11 in which a heat exchange fluid
10 circulates. The fluid leaving the recipient 5 is thus taken to a state of desired pressure and temperature determined during its passage in the exchanger 7 and the pressure reducing valve 8, so that it comprises a gaseous phase and a liquid phase. The heating means constituted by the double envelope 11 make it possible to vaporize liquid phase arriving in the impregnation enclosure 9, so as to
15 maintain constant the level of liquid phase therein. Under these conditions, a gaseous phase is therefore had permanently in the impregnation enclosure 9, constituted by solvent fluid and a liquid phase constituted by the extracts and the solvent fluid.

 The porous medium 10 is chosen as a function of the subsequent use which
20 it is desired to make of the final product, whether it be a granular or pulverulent medium particularly adapted for use in dietetics, pharmacy or cosmetics, or a solid medium. In the most frequent case where this medium is in granular or pulverulent form, a particularly advantageous embodiment consists in stirring this porous medium within the impregnation enclosure 9 by any

The upper part of the enclosure 9 is joined to the pump 3 through a condenser 12.

Of course, it is possible, according to the invention, to use any other separation system, and the extractor 1 might be replaced by a fractionating column making possible

the continuous or discontinuous treatment of raw materials in the liquid state, the fluid emerging from the column being treated in the same manner as that described previously.

In a variant of the invention, an auxiliary pump is used for introducing in the extractor 1 one or more organic co-solvents making it possible to modify the solvent power and the polarity of the solvent fluid. It is often chosen to add ethanol which may be of alimentary quality or CODEX depending on the destination of the products thus elaborated. A light hydrocarbon having between 2 and 8 carbon atoms may also favourably be used as co-solvent. In the case of a co-solvent being used, a porous medium which is not altered by this co-solvent will naturally be chosen. Moreover, at the end of the extraction-impregnation operation, care will be taken to scavenge the porous medium with pure solvent fluid without co-solvent in order to eliminate the adsorbed co-solvent.

It has been ascertained that, under these conditions, an excellent diffusion of the fluid was obtained through the pores of the porous medium, and, correlatively, the entrainment in these latter of the extracts and their adsorption in the pores.

A contribution of enthalpy is required to maintain a constant quantity of fluid in the liquid state within the impregnation enclosure 9, in proportion as the fluid coming from extraction is injected and as an identical delivery of fluid which emerges from this enclosure in the gaseous state is therefore vaporized. This contribution of enthalpy must be carefully adjusted in order to avoid, if it is insufficient, either the accumulation of liquefied fluid in the

impregnation recipient which would finish by emerging in this liquid form, taking along part of the extracts, or, if it is too great, the total vaporization of the fluid and the non-controlled precipitation of the extracts.

As will be illustrated in the following examples of implementation of the method and of the installation according to the invention, it is surprising to effect extraction and to obtain a very homogeneous impregnation of different excipients by the extracts in one operation, without ever having to manipulate the extracts themselves, which can but avoid all risk of degradation by oxidation in air or by exposure to heat, since any contact with air is avoided and all the operations are conducted at a temperature close to ambient temperature.

Several examples of the invention which were carried out with the installation described hereinabove and whose operational parameters were as follows, will be described hereinafter:

Volume of the basket 2	: 0.5 l
Flowrate of the pump 3	: 1 to 5 kg/hr
Working pressure	: of the order of 30 MPa
Working temperature	: between 10°C and 80°C
Contents of recipient 5	: 100 g of zeolite 3A
Volume of impregnation enclosure 9	: 4.5 l
Porous material	: Maltodextrine powder of alimentary quality obtained by partial hydrolysis of maize amide
Speed of rotation of the turbine 18	: 120 revs/minute

Example 1: Extraction and impregnation of KAVA-KAVA:

Kava-kava is the local name of a wild shrub of the Pacific islands, identified as *Piper methysticum* or *Piper wichmannu*, whose roots contain products of great pharmacological interest called kavalactones, widely used in different forms as natural tranquillizer and euphoriant.

The extraction was conducted on 100 g of powder of dried roots, ground towards about 200 μm , with a flowrate of 3 kg/hr of carbon dioxide at 25 MPa and 40°C, and furnished an extract which is in the form of a very viscous, dark yellow paste with characteristic odour. After adsorption of the water, the pressure of the fluid was reduced to 6 MPa and injected via the bottom into the impregnation enclosure 9 containing 100 g of maltodextrine. It was observed that the compressed gas leaving the top of the enclosure 9 presented a constant temperature close to 40°C in established operation. After having continued the operation for 510 mins., the impregnation enclosure 9 was decompressed and 114.8 g of a bright yellow powder was recovered therein, which presented the characteristic odour of the extract of kava-kava and flowing without problem in the absence of any lump or agglomerate, ideal for manufacturing tablets, possibly mixed with an excipient or other active principles. A sample of this powder was re-extracted with chloroform and analyzed by gas-chromatography. It was ascertained that the extract fixed on the maltodextrine was constituted by 89% by mass of kavalactones whose identification makes it possible to verify that the relative abundance of each of these compounds is conforming

to what is found in a conventional extract. This therefore confirms both the high selectivity of the extraction by the carbon dioxide at supercritical pressure and the complete fixation of the extract on the maltodextrine.

Example 2: Extraction and impregnation of KAVA-KAVA:

5 A second operation was proceeded with under conditions identical to those used in Example 1, except that, this time, the initial mass of maltodextrine was only 50 g. After 510 mins. of operation, 65.6 g of powder were obtained, whose characteristics are identical to those of the powder obtained in Example 1, except that its colour and odour are more intense. An analysis of the fixed extract
10 revealed a percentage of kavalactones of 91% by mass, with a distribution between the different compounds virtually identical to that observed in the preceding Example. This shows that the maltodextrine may be charged with at least 30% by mass of extract.

Example 3: Extraction and impregnation of KAVA-KAVA:

15 A third operation was proceeded with under conditions identical to those used in Example 2, except that, this time, a porous medium was chosen, composed of an intimate mixture of powders of maltodextrine and soja lecithin of alimentary quality, at a rate of 45 g of maltodextrine for 5 g of lecithin. The initial mass of porous medium was fixed at 50 g. After 510 mins. of operation, 64.2 g of
20 powder were obtained, whose characteristics proved to be close to those of the powder obtained in Example 1, except that its

colour and odour are more intense. An analysis of the fixed extract led to a percentage of kavalactones of 90% by mass, with a distribution between the different compounds virtually identical to that observed in the preceding Example. This powder thus presents the advantage, with respect to the powders
5 obtained in Examples 1 and 2, of being more easily dispersed in water, giving rise to a turbidity resembling that obtained by diluting pastis in water. It may therefore be used not only in dry formulations, such as tablets, but also in the form of a powder to be mixed in water in order to prepare a drinkable potion.

Example 4: Extraction and impregnation of a spice (Curcuma):

10 The extraction was conducted on a total mass of 800 g of ground Curcuma, distributed in 4 batches of 200 g placed successively in the extractor 1 with a flowrate of 2.4 kg/hr of carbon dioxide at 40°C and 29 MPa. After elimination of the water, the pressure of the latter was reduced to 5 MPa and it was injected via the bottom in the impregnation enclosure 9 containing a mass of 400 g of
15 maltodextrine which was successively impregnated with the extracts issuing from the four batches. After having continued the operation for four periods of 50 mins. each, 445 g of a very homogeneous orange powder were recovered, presenting the odour and characteristic taste of the extract of Curcuma.

Example 5: Extraction and impregnation of a mixture of spices (black
20 pepper + mild paprika):

One proceeded with an operation of extraction and impregnation of the spices under conditions identical to

those used in Example 4, except that, this time, the charge was constituted by a mixture of 180 g of pepper and 20 g of paprika. After having passed 8.4 kg of carbon dioxide, 412 g of very homogeneous red powder were obtained, in the absence of lump or agglomerate.

	1970	1971	1972	1973	1974	1975	1976	1977	1978	1979	1980	1981	1982	1983	1984	1985	1986	1987	1988	1989	1990	1991	1992	1993	1994	1995	1996	1997	1998	1999	2000	2001	2002	2003	2004	2005	2006	2007	2008	2009	2010	2011	2012	2013	2014	2015	2016	2017	2018	2019	2020	2021	2022	2023	2024	2025	2026	2027	2028	2029	2030	2031	2032	2033	2034	2035	2036	2037	2038	2039	2040	2041	2042	2043	2044	2045	2046	2047	2048	2049	2050	2051	2052	2053	2054	2055	2056	2057	2058	2059	2060	2061	2062	2063	2064	2065	2066	2067	2068	2069	2070	2071	2072	2073	2074	2075	2076	2077	2078	2079	2080	2081	2082	2083	2084	2085	2086	2087	2088	2089	2090	2091	2092	2093	2094	2095	2096	2097	2098	2099	2100	2101	2102	2103	2104	2105	2106	2107	2108	2109	2110	2111	2112	2113	2114	2115	2116	2117	2118	2119	2120	2121	2122	2123	2124	2125	2126	2127	2128	2129	2130	2131	2132	2133	2134	2135	2136	2137	2138	2139	2140	2141	2142	2143	2144	2145	2146	2147	2148	2149	2150	2151	2152	2153	2154	2155	2156	2157	2158	2159	2160	2161	2162	2163	2164	2165	2166	2167	2168	2169	2170	2171	2172	2173	2174	2175	2176	2177	2178	2179	2180	2181	2182	2183	2184	2185	2186	2187	2188	2189	2190	2191	2192	2193	2194	2195	2196	2197	2198	2199	2200	2201	2202	2203	2204	2205	2206	2207	2208	2209	2210	2211	2212	2213	2214	2215	2216	2217	2218	2219	2220	2221	2222	2223	2224	2225	2226	2227	2228	2229	2230	2231	2232	2233	2234	2235	2236	2237	2238	2239	2240	2241	2242	2243	2244	2245	2246	2247	2248	2249	2250	2251	2252	2253	2254	2255	2256	2257	2258	2259	2260	2261	2262	2263	2264	2265	2266	2267	2268	2269	2270	2271	2272	2273	2274	2275	2276	2277	2278	2279	2280	2281	2282	2283	2284	2285	2286	2287	2288	2289	2290	2291	2292	2293	2294	2295	2296	2297	2298	2299	2300	2301	2302	2303	2304	2305	2306	2307	2308	2309	2310	2311	2312	2313	2314	2315	2316	2317	2318	2319	2320	2321	2322	2323	2324	2325	2326	2327	2328	2329	2330	2331	2332	2333	2334	2335	2336	2337	2338	2339	2340	2341	2342	2343	2344	2345	2346	2347	2348	2349	2350	2351	2352	2353	2354	2355	2356	2357	2358	2359	2360	2361	2362	2363	2364	2365	2366	2367	2368	2369	2370	2371	2372	2373	2374	2375	2376	2377	2378	2379	2380	2381	2382	2383	2384	2385	2386	2387	2388	2389	2390	2391	2392	2393	2394	2395	2396	2397	2398	2399	2400	2401	2402	2403	2404	2405	2406	2407	2408	2409	2410	2411	2412	2413	2414	2415	2416	2417	2418	2419	2420	2421	2422	2
--	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	------	---

CLAIMS

1. Method for setting in adsorbed state, on a porous support, compounds
5 contained in a product, in which, during a first step, the extraction of the
compounds is effected by contacting the product with at least one solvent fluid at
supercritical pressure leading to the obtaining of a mixture of extracts and of
solvent fluid, and, in a second step:

- the water contained in the mixture of extracts and of solvent fluid is
10 eliminated, by causing the latter to trickle over a bed of adsorbent product
adapted to fix this water selectively,

- the temperature and pressure conditions are adjusted so as to obtain, in an
enclosure (9), two phases, namely a first phase essentially constituted by the
solvent fluid in the gaseous state and a second phase constituted by a mixture of
15 liquids formed by solvent fluid and extracts of the product,

- these two phases are made to trickle through a porous support adapted to
adsorb the extracts,

- the solvent fluid contained in the second phase is vaporized.

2. Method according to Claim 1, characterized in that the solvent fluid is
20 constituted by carbon dioxide, by nitrogen protoxide or by a light hydrocarbon
having from 2 to 8 carbon atoms.

3. Method according to one of the preceding Claims, characterized in that the
solvent fluid is a

AMENDED SHEET

pure fluid.

4. Method according to one of Claim 1 or 2, characterized in that the solvent
5 fluid has at least one co-solvent added thereto.

5. Method according to Claim 4, characterized in that the co-solvent is
constituted by an alcohol and preferably ethanol, and/or by a ketone and
preferably acetone, and/or by an ester and preferably ethyl acetate.

6. Method according to one of the preceding Claims, characterized in that the
10 first step of extraction is effected at a pressure included between 7.4 MPa and 80
MPa, and preferably between 10 MPa and 40 MPa, and at a temperature included
between 0°C and 80°C.

7. Method according to one of the preceding Claims, characterized in that the
15 trickling of the two phases through the porous support is effected at a pressure
included between 1 MPa and 10 MPa, and preferably between 4 MPa and 8 Mpa,
and at a temperature included between 0°C and 80°C.

8. Installation for extraction/impregnation of the type comprising an extractor
(1) and an impregnation enclosure (9) containing a porous medium, the extractor
(1) containing a product from which it is desired to extract the compounds, which
20 is traversed to that end by at least one solvent fluid at supercritical pressure,
characterized in that it successively comprises, downstream of the extractor (1),
means (5) for eliminating the water contained in the extracted compounds, means
(11) for contributing enthalpy, adapted to create, in the impregnation enclosure
(9), two phases, namely a

first phase essentially constituted by the solvent fluid in the gaseous state and a second phase constituted by a mixture of liquids formed by solvent fluid and the extracts of the product, so as to effect the adsorption by the porous medium of the extracted compounds.

9. Installation according to Claim 8, characterized in that the means for contributing enthalpy are constituted by a double envelope (11) with circulation of heat-exchange fluid.

10. Installation according to one of Claims 8 or 9, characterized in that the extractor (1) is constituted by a fractionating column operating in counter-flow, adapted for the treatment of liquid raw materials.

11. Installation according to one of Claims 8 to 10, characterized in that it comprises means for injecting an organic co-solvent within the solvent fluid.

AMENDED SHEET

METHOD AND INSTALLATION FOR SETTING IN ADSORBED STATE
ON A POROUS SUPPORT ACTIVE COMPOUNDS CONTAINED IN A
PRODUCT

5

The invention concerns a method and an installation for setting in adsorbed state, on a porous support, compounds contained in a product comprising a first step which consists in extracting the compounds by contacting the product with at least a solvent at supercritical pressure to obtain a mixture of extracts and solvent.

10

The method is characterized in that it comprises a second step which consists in eliminating the water contained in the mixture of extracts and solvent, adjusting the temperature and pressure conditions so as to obtain, a first phase consisting of solvent in gaseous state and a second phase consisting of a mixture of liquids formed of solvent and extracts of the products; causing said two phases to trickle through a porous support adapted to adsorb the extracts; vaporizing the solvent contained in the second phase.

15

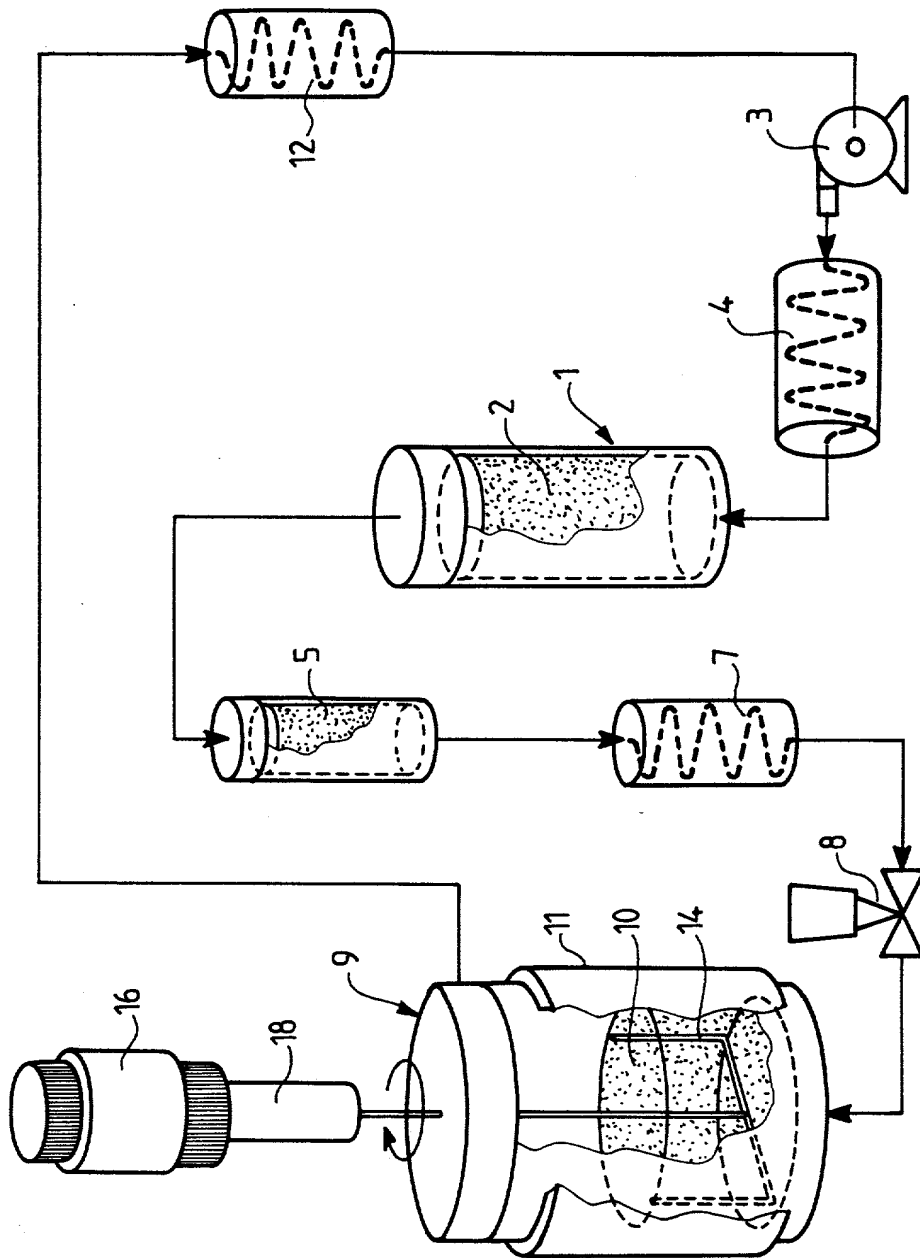


FIGURE UNIQUE

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it contains a valid OMB control number.

**DECLARATION FOR UTILITY OR
DESIGN
PATENT APPLICATION
(37 CFR 1.63)**

☒ Declaration Submitted with Initial Filing OR ☐ Declaration Submitted after Initial Filing (surcharge (37 CFR 1.16 (e)) required)

Attorney Docket Number

366325-525

First Named Inventor

Perrut, Michel

COMPLETE IF KNOWN

Application Number

/

Filing Date

Art Unit

Examiner Name

As the below named inventor, I hereby declare that:

My residence, mailing address, and citizenship are as stated below next to my name.

I believe I am the original and first inventor of the subject matter which is claimed and for which a patent is sought on the invention entitled:

**METHOD AND INSTALLATION FOR SETTING IN ADSORBED STATE ON A
POROUS SUPPORT ACTIVE COMPOUNDS CONTAINED IN A PRODUCT**

(Title of the Invention)

the specification of which

☒ is attached hereto

OR

☐ was filed on (MM/DD/YYYY)

as United States Application Number or PCT International

Application Number

and was amended on (MM/DD/YYYY)

(if applicable).

I hereby state that I have reviewed and understand the contents of the above identified specification, including the claims, as amended by any amendment specifically referred to above.

I acknowledge the duty to disclose information which is material to patentability as defined in 37 CFR 1.56, including for continuation-in-part applications, material information which became available between the filing date of the prior application and the national or PCT international filing date of the continuation-in-part application.

I hereby claim foreign priority benefits under 35 U.S.C. 119(a)-(d) or (f), or 365(b) of any foreign application(s) for patent, inventor's or plant breeder's rights certificate(s), or 365(a) of any PCT international application which designated at least one country other than the United States of America, listed below and have also identified below, by checking the box, any foreign application for patent, inventor's or plant breeder's rights certificate(s), or any PCT international application having a filing date before that of the application on which priority is claimed.

Prior Foreign Application Number(s)	Country	Foreign Filing Date (MM/DD/YYYY)	Priority Not Claimed	Certified Copy Attached?	
				YES	NO
99/12005	FR	09/27/1999	<input type="checkbox"/>	<input type="checkbox"/>	<input checked="" type="checkbox"/>
			<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
			<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
			<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>

☐ Additional foreign application numbers are listed on a supplemental priority data sheet PTO/SB/02B attached hereto:

[Page 1 of 2]

PTO/SB/01 (10-01)

Approved for use through 10/31/2002. OMB 0651-0032

U.S. Patent and Trademark Office: U.S. DEPARTMENT OF COMMERCE

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it contains a valid OMB control number.

DECLARATION — Utility or Design Patent ApplicationDirect all correspondence to: ☒ Customer Number 25561 OR ☐ Correspondence address below
or Bar Code Label

Name

Address

City

State

ZIP

Country

Telephone

Fax

I hereby declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under 18 U.S.C. 1001 and that such willful false statements may jeopardize the validity of the application or any patent issued thereon.

NAME OF SOLE OR FIRST INVENTOR : ☐ A petition has been filed for this unsigned inventorGiven Name
(first and middle [if any])MichelFamily Name
or SurnamePerrutInventor's
SignatureMichel Perrut

Date

14.03.02

Residence: City

Nancy

State

Country

France

Citizenship

French

Mailing Address

25, rue de Santifontaine

City

54000 Nancy

State

ZIP

Country

France

NAME OF SECOND INVENTOR:

☐ A petition has been filed for this unsigned inventorGiven Name
(first and middle [if any])WieslawFamily Name
or SurnameMajewskiInventor's
SignatureWieslaw Majewski

Date

Residence: City

Laxou

State

Country

FR

Citizenship

FR

Mailing Address

"Le Meridien", 4, terrasse des Vosges

City

54520 Laxou

State

ZIP

Country

FR☐ Additional inventors are being named on the _____ supplemental Additional Inventor(s) sheet(s) PTO/SB/02A attached hereto.